

L Number	Hits	Search Text	DB	Time stamp
3	658	BIR\$5 WITH domain	USPAT; US-PGPUB; EPO; JPO; DERWENT	2004/06/10 14:05
5	78	(BIR\$5 WITH domain) and iap	USPAT; US-PGPUB; EPO; JPO; DERWENT	2004/06/10 14:05
6	51	(XIAP M-XIAP HIAP\$3 M-HIAP\$3) SAME BIR\$3	USPAT; US-PGPUB; EPO; JPO; DERWENT	2004/06/10 14:05
7	14	(US-6511828-\$ or US-6495339-\$ or US-6472172-\$ or US-6331412-\$ or US-6300492-\$ or US-6228603-\$ or US-6187557-\$ or US-6171821-\$ or US-6159709-\$ or US-6156535-\$ or US-6133437-\$ or US-6107088-\$ or US-6107041-\$ or US-6087173-\$ or US-5919912-\$).did. or (US-20020120121-\$ or US-20020086409-\$ or US-20020187946-\$ or US-20020160975-\$ or US-20020132786-\$ or US-20020137028-\$).did. or (WO-9706255-\$ or EP-892048-\$ or WO-9835693-\$ or WO-9822131-\$ or WO-9740847-\$ or WO-9726331-\$ or WO-9612016-\$ or WO-9316196-\$).did. or (JP-11032780-\$).did.	USPAT; US-PGPUB; EPO; JPO	2004/06/10 14:05
8	15	(BIR-3 OR BIR3) SAME apoptosis	USPAT; US-PGPUB; EPO; JPO; DERWENT	2004/06/10 14:10
9	37	Robert WITH KORNELUK,	USPAT; US-PGPUB; EPO; JPO; DERWENT	2004/06/10 14:13
10	34	(US-6511828-\$ or US-6472172-\$ or US-6228603-\$ or US-6187557-\$ or US-6087173-\$ or US-6331412-\$ or US-6495339-\$ or US-6300492-\$ or US-6171821-\$ or US-6159709-\$ or US-6156535-\$ or US-6133437-\$ or US-6107088-\$ or US-6107041-\$ or US-5919912-\$ or US-6656704-\$ or US-6689562-\$).did. or (US-20020160975-\$ or US-20020132786-\$ or US-20020120121-\$ or US-20020137028-\$ or US-20020086409-\$ or US-20020187946-\$ or US-20030157522-\$).did. or (WO-9706255-\$ or EP-892048-\$ or WO-9835693-\$ or WO-9822131-\$ or WO-9726331-\$ or WO-9740847-\$ or WO-9316196-\$ or WO-9612016-\$).did. or (JP-11032780-\$).did. or (WO-200216418-\$).did.	USPAT; US-PGPUB; EPO; JPO; DERWENT	2004/06/10 14:13

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(FILE 'HOME' ENTERED AT 14:16:07 ON 10 JUN 2004)

FILE 'MEDLINE' ENTERED AT 14:16:23 ON 10 JUN 2004

L1 24 S (BIR-3 OR BIR3) (L) APOPTOSIS

FILE 'MEDLINE, SCISEARCH, CAPLUS, MEDICONF' ENTERED AT 14:17:38 ON 10 JUN 2004

L2 85 S L1

L3 37 DUP REM L2 (48 DUPLICATES REMOVED)

L4 37 SORT L3 PY

E KOMELUK ROBERT?/AU

L5 3 S E1

L6 1 S E2

L7 4 S L5 OR L6

L8 37 FOCUS L4 1-

=> d an ti so au ab 18 6

L8 ANSWER 6 OF 37 MEDLINE on STN

AN 1999438002 MEDLINE

TI Cleavage of human inhibitor of apoptosis protein XIAP results in fragments with distinct specificities for caspases.

SO EMBO journal, (1999 Oct 1) 18 (19) 5242-51.

Journal code: 8208664. ISSN: 0261-4189.

AU Deveraux Q L; Leo E; Stennicke H R; Welsh K; Salvesen G S; Reed J C

AB Several human inhibitor of **apoptosis** (IAP) family proteins function by directly inhibiting specific caspases in a mechanism that does not require IAP cleavage. In this study, however, we demonstrate that endogenous XIAP is cleaved into two fragments during **apoptosis** induced by the tumor necrosis factor family member Fas (CD95). The two fragments produced comprise the baculoviral inhibitory repeat (BIR) 1 and 2 domains (BIR1-2) and the **BIR3** and RING (**BIR3**-Ring) domains of XIAP. Overexpression of the BIR1-2 fragment inhibits Fas-induced **apoptosis**, albeit at significantly reduced efficiency compared with full-length XIAP. In contrast, overexpression of the **BIR3**-Ring fragment results in a slight enhancement of Fas-directed **apoptosis**. Thus, cleavage of XIAP may be one mechanism by which cell death programs circumvent the anti-apoptotic barrier posed by XIAP. Interestingly, ectopic expression of the **BIR3**-Ring fragment resulted in nearly complete protection from Bax-induced **apoptosis**. Use of purified recombinant proteins revealed that **BIR3**-Ring is a specific inhibitor of caspase-9 whereas BIR1-2 is specific for caspases 3 and 7. Therefore XIAP possesses two different caspase inhibitory activities which can be attributed to distinct domains within XIAP. These data may provide an explanation for why IAPs have evolved with multiple BIR domains.

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